

**Amendments to the Claims:**

The listing of claims will replace all prior versions and listings, of claims in the application:

**Listing of Claims:**

Claims 1-5 (canceled)

Claim 6 (currently amended) A method of determining the relative level of *Dihydropyrimidine dehydrogenase (DPD)* gene expression in a tissue sample comprising:

- (a) obtaining a tumor sample from a patient;
- (b) fixing at least a portion of said tumor sample in paraffin to achieve a fixed and paraffin embedded (FPE) tumor tissue sample;
- (c) isolating mRNA from said (FPE) tumor tissue sample;
- (d) determining the amount of *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from said FPE tumor tissue sample by amplifying the mRNA using an oligonucleotide primer SEQ ID: 1, ~~or an oligonucleotide primer at least 80% identical therewith and hybridizes to a complement of SEQ ID NO. 1 under stringent conditions; wherein said isolated and purified oligonucleotide is capable of amplifying a portion of the 5' untranslated region and Exon 1 of a *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from fixed and paraffin embedded (FPE) tissue when used with SEQ ID NO. 2; and; oligonucleotide primer SEQ ID: 2 or an oligonucleotide primer at least 80% identical therewith and hybridizes to a complement of SEQ ID NO. 2 under stringent conditions; wherein said isolated and purified oligonucleotide is capable of amplifying a portion of the 5' untranslated region and Exon 1 of a *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from fixed and paraffin embedded (FPE) tissue when used with SEQ ID NO. 1;~~
- (e) comparing the amount of *Dihydropyrimidine dehydrogenase (DPD)* mRNA from step (d) to an amount of mRNA of an internal control gene.

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Claims 7-9 (canceled)

Claim 10 (currently amended) The method of ~~claim 8 or 9~~ claim 6, wherein the RNA is isolated in the presence of an effective amount of chaotropic agent.

Claim 11 (currently amended) The method of ~~any one of claims~~ claim 6, 8, or 9, wherein the tumor sample comprises non-tumor tissue and tumor tissue.

Claims 12-16 (canceled)

Claim 17 (currently amended) A method of determining the relative level of *Dihydropyrimidine dehydrogenase (DPD)* gene expression in a tissue sample comprising;

- (a) obtaining a tumor sample from a patient;
- (b) fixing at least a portion of said tumor sample in paraffin to achieve a fixed and paraffin embedded (FPE) tumor tissue sample;
- (b)(c) isolating mRNA from said FPE tumor tissue sample;
- (c)(d) determining the amount of *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from said FPE tumor tissue sample by amplifying the mRNA using an oligonucleotide primer SEQ ID: 7, ~~or an oligonucleotide primer at least 80% identical therewith and hybridizes to a complement of SEQ ID NO: 7 under stringent conditions, wherein said isolated and purified oligonucleotide is capable of amplifying a portion of Exon 6 of a *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from fixed and paraffin embedded (FPE) tissue when used with SEQ ID NO: 8; and; oligonucleotide primer SEQ ID: 8 or an oligonucleotide primer at least 80% identical therewith and hybridizes to a complement of SEQ ID NO: 8 under stringent conditions, wherein said isolated and purified oligonucleotide is capable of amplifying a portion of Exon 6 of a~~

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*Dihydropyrimidine dehydrogenase (DPD) mRNA isolated from fixed and paraffin embedded (FPE) tissue when used with SEQ ID NO: 7;*

- (d)(e) comparing the amount of the mRNA from step (c) (d) to an amount of mRNA of an internal control.

Claims 18-19 (canceled)

Claim 20 (currently amended) The method of claim 19 claim 17, wherein the mRNA isolated from said FPE tumor tissue is isolated in the presence of an effective amount of chaotropic agent.

Claim 21 (canceled)

Claim 22 (currently amended) The method of claim 20 claim 17, wherein a FPE tumor tissue sample comprises non-tumor tissue and tumor tissue.

Claims 23-26 (canceled)

Claim 27 (currently amended) A method of determining the relative level of *Dihydropyrimidine dehydrogenase (DPD)* gene expression in a tissue sample comprising:

- (a) obtaining a tumor sample from a patient, wherein said tumor sample is fixed and paraffin embedded (FPE);
- (b) isolating mRNA from said FPE tumor sample, wherein said FPE tumor sample is heated to a temperature in the range of about 50 to about 100°C;
- (c) determining the amount of *Dihydropyrimidine dehydrogenase (DPD)* mRNA by amplifying the mRNA using an oligonucleotide primer SEQ ID: 1, or an oligonucleotide primer at least 80% identical therewith and hybridizes to a complement of SEQ ID NO: 1 under stringent conditions, wherein said isolated

- and purified oligonucleotide is capable of amplifying a portion of the 5' untranslated region and Exon 1 of a *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from fixed and paraffin embedded (FPE) tissue when used with SEQ ID NO: 2; and; oligonucleotide primer SEQ ID: 2 or an oligonucleotide primer at least 80% identical therewith and hybridizes to a complement of SEQ ID NO: 2 under stringent conditions; wherein said isolated and purified oligonucleotide is capable of amplifying a portion of the 5' untranslated region and Exon 1 of a *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from fixed and paraffin embedded (FPE) tissue when used with SEQ ID NO: 1;
- (d) comparing the amount of *Dihydropyrimidine dehydrogenase (DPD)* mRNA from step (c) to an amount of mRNA of an internal control gene.

Claim 28 (currently amended) A method of determining the relative level of *Dihydropyrimidine dehydrogenase (DPD)* gene expression in a tissue sample comprising:

- (a) obtaining a tumor sample from a patient, wherein said tumor sample is fixed and paraffin embedded (FPE);
- (b) isolating mRNA from said FPE tumor sample, wherein said FPE tumor sample is heated to a temperature in the range of about 50 to about 100°C;
- (c) determining the amount of *Dihydropyrimidine dehydrogenase (DPD)* mRNA by amplifying the mRNA using an oligonucleotide primer SEQ ID: 7, or an oligonucleotide primer at least 80% identical therewith and hybridizes to a complement of SEQ ID NO: 7 under stringent conditions; wherein said isolated and purified oligonucleotide is capable of amplifying a portion of Exon 6 of a *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from fixed and paraffin embedded (FPE) tissue when used with SEQ ID NO: 8; and; oligonucleotide primer SEQ ID: 8 or an oligonucleotide primer at least 80% identical therewith and hybridizes to a complement of SEQ ID NO: 8 under stringent conditions;

- ~~wherein said isolated and purified oligonucleotide is capable of amplifying a portion of the 5' untranslated region and Exon 6 of a *Dihydropyrimidine dehydrogenase (DPD)* mRNA isolated from fixed and paraffin embedded (FPE) tissue when used with SEQ ID NO. 7;~~
- (d) comparing the amount of *Dihydropyrimidine dehydrogenase (DPD)* mRNA from step (c) to an amount of mRNA of an internal control gene.

Claim 29 (previously presented) The method of claims 27 or 28, wherein said internal control gene is  $\beta$ -actin.

Claim 30 (canceled)

Claim 31 (previously presented) The method of claims 27 or 28, wherein the mRNA is isolated in the presence of an effective amount of chaotropic agent.

Claim 32 (currently amended) A method for determining the relative level of *Dihydropyrimidine dehydrogenase (DPD)* gene expression in a fixed and paraffin embedded (FPE) tissue sample comprising:

- (a) deparaffinizing the FPE tissue sample to obtain a deparaffinized sample;
- (b) isolating mRNA from the deparaffinized sample in the presence of an effective amount of a chaotropic agent by first heating the tissue sample in a solution comprising an effective concentration of a chaotropic chaotropic compound to a temperature in the range of about 75° to about 100° C for a time period of 5 to 120 minutes and recovering said mRNA from said chaotropic solution; and
- (c) subjecting the mRNA to amplification using a pair of oligonucleotide primers capable of amplifying a region of the ERCC1 DPD gene, to obtain an amplified sample; wherein the pair of primers is primer pair SEQ ID NO:1 and 2 or primer

pair SEQ ID NO 7 and 8;

- (d) determining the quantity of ~~ERCC1 DPD~~ mRNA relative to the quantity of an internal control gene's mRNA.

Claim 33 (previously presented) The method of claim 32 wherein, the internal control gene is  $\beta$ -actin.

Claim 34 (currently amended) A method for determining the relative level of *Dihydropyrimidine dehydrogenase* (DPD) gene expression in a fixed and paraffin embedded tissue sample comprising:

- (a) deparaffinizing the tissue sample to obtain a deparaffinized sample;
- (b) isolating mRNA from the deparaffinized sample by first heating the deparaffinized tissue sample in a solution comprising an effective concentration of a chaotropic agent to a temperature in the range of about 50° to about 100° C and recovering said mRNA from said solution; and
- (c) determining the quantity of ~~ERCC1 DPD~~ mRNA relative to the quantity of an internal control gene's mRNA.

Claim 35 (currently amended) A method for determining a platinum-based chemotherapeutic regimen for treating a tumor in a patient comprising:

- (a) obtaining a tissue sample of the tumor and fixing the sample to obtain a fixed tumor sample;
- (b) isolating mRNA from the fixed tumor sample, wherein the fixed tumor sample is heated in the presence of an effective amount of a chaotropic chaotropic agent and wherein the heating occurs at a temperature from about 50° to about 100° C;
- (c) subjecting the mRNA to amplification using a pair of oligonucleotide primers capable of amplifying a region of the *Dihydropyrimidine dehydrogenase* (DPD)

- gene to obtain an amplified sample; wherein the pair of primers is primer pair SEQ ID NO:1 and 2 or primer pair SEQ ID NO 7 and 8;
- (d) determining the amount of amplified DPD mRNA in the amplified sample;
  - (e) comparing the amount of DPD mRNA from step (d) to an amount of mRNA of an internal control gene; and
  - (f) determining a platinum-based chemotherapeutic regimen based on the amount of DPD mRNA in the amplified sample and a predetermined threshold level for DPD gene expression.

Claim 36 (canceled)

Claim 37 (currently amended) The method of claim 35 wherein the fixed tumor sample is heated in the presence of an effective amount of a ~~chaotropic~~ chaotropic agent and wherein the heating occurs at a temperature from about 75 °C to about 100 °C for a period of about 5 to about 120 minutes.